

region of **adenovirus 5**. This construct directed high levels of synthesis of **p53** in HeLa cells.

L9 ANSWER 42 OF 42 MEDLINE on STN DUPLICATE 15  
ACCESSION NUMBER: 95134785 MEDLINE  
DOCUMENT NUMBER: PubMed ID: 7833367  
TITLE: Development and characterization of recombinant **adenoviruses** encoding human **p53** for **gene therapy** of cancer.  
AUTHOR: Wills K N; Maneval D C; Menzel P; Harris M P; Sutjipto S; Vaillancourt M T; Huang W M; Johnson D E; Anderson S C; Wen S F; +  
CORPORATE SOURCE: CANJI, Inc., San Diego, CA 92121.  
SOURCE: Human gene therapy, (1994 Sep) 5 (9) 1079-88.  
Journal code: 9008950. ISSN: 1043-0342.  
PUB. COUNTRY: United States  
DOCUMENT TYPE: Journal; Article; (JOURNAL ARTICLE)  
LANGUAGE: English  
FILE SEGMENT: Priority Journals  
ENTRY MONTH: 199502  
ENTRY DATE: Entered STN: 19950314  
Last Updated on STN: 19970203  
Entered Medline: 19950224

AB We have constructed recombinant human **adenoviruses** that express wild-type human **p53** under the control of either the Ad 2 major late promoter (MLP) or the human cytomegalovirus (**CMV**) immediate early gene promoter. Each construct replaces the Ad 5 Ela and Elb coding sequences necessary for viral replication with the **p53** cDNA and MLP or **CMV** promoter. These **p53**/Ad recombinants are able to express **p53** protein in a dose-dependent manner in infected human cancer cells. Tumor suppressor activity of the expressed **p53** protein was assayed by several methods. [3H]Thymidine incorporation assays showed that the recombinant **adenoviruses** were capable of inhibiting DNA synthesis in a **p53**-specific, dose-dependent fashion. Ex vivo treatment of Saos-2 tumor cells, followed by injection of the treated cells into nude mice, led to complete tumor suppression using the MLP/**p53** recombinant. Following a single injection of **CMV/p53** recombinant **adenovirus** into the peritumoral space surrounding an in vivo established tumor derived from a human small cell lung carcinoma cell line (NIH-H69), we were able to detect **p53** mRNA in the tumors at 2 and 7 days post-injection. Continued treatment of established H69 tumors with MLP/**p53** recombinant led to reduced tumor growth and increased survival time compared to control treated animals. These results indicate that recombinant **adenoviruses** expressing wild-type **p53** may be useful vectors for **gene therapy** of human cancer.


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FILE 'MEDLINE' ENTERED AT 06:24:50 ON 28 SEP 2004

L1 29300 S ADENOVIR?  
L2 681 S L1 AND (CMV OR CMV?)  
L3 33 S L2 AND P53  
L4 22 S L3 AND CANCER  
L5 24 S L3 AND NEOPLASM  
L6 14 S L5 AND TREATMENT  
L7 19 S L5 AND GENE(W) THERAPY

FILE 'MEDLINE, BIOSIS, EMBASE, CAPLUS, SCISEARCH, WPIDS' ENTERED AT

 Files searched

06:26:11 ON 28 SEP 2004

L8

58 S L7

L9

42 DUP REM L8 (16 DUPLICATES REMOVED)

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L5

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| <u>L1</u> | adenovir\$ and p53 and (cmv or cmv\$) | 3052 | <u>L1</u> |

END OF SEARCH HISTORY